

[0096] 54. A cell culture system comprising:  
[0097] a. a host cell capable of producing rAAV;  
[0098] b. a helper virus; and  
[0099] c. a cell culture medium with an osmolality of 360 mOsm/kg or higher when measured immediately after the host cell is introduced into the cell culture medium.  
[0100] 55. The cell culture system of claim 54, wherein the cell culture medium has an osmolality of 375 mOsm/kg or higher.  
[0101] 56. The cell culture system of claim 55, wherein the cell culture medium has an osmolality of 400 mOsm/kg or higher.  
[0102] 57. The cell culture system of any one of claims 54-56, wherein the cell culture medium has an osmolality sufficient to produce a 20% reduction in total helper virus production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0103] 58. The cell culture system of any one of claims 54-56, wherein the cell culture medium has an osmolality sufficient to produce a 30% reduction in total helper virus production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0104] 59. The cell culture system of any one of claims 54-56, wherein the cell culture medium has an osmolality sufficient to produce a 40% reduction in total helper virus production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0105] 60. The cell culture system of any one of claims 54-56, wherein the cell culture medium has an osmolality sufficient to produce a 50% reduction in total helper virus production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0106] 61. The cell culture system of any one of claims 54-56, wherein the cell culture medium has an osmolality sufficient to produce at least a 50% increase in total rAAV production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0107] 62. The cell culture system of any one of claims 54-56, wherein the cell culture medium has an osmolality sufficient to produce at least a 100% increase in total rAAV production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0108] 63. The cell culture system of any one of claims 54-56, wherein the cell culture medium has an osmolality sufficient to produce at least a 150% increase in total rAAV production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0109] 64. The cell culture system of any one of claims 54-56, wherein the cell culture medium has an osmolality sufficient to produce at least a 200% increase in total rAAV production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0110] 65. The cell culture system of any one of claims 54-64, wherein at least one tonicifying agent is an ionic tonicifying agent.  
[0111] 66. The cell culture system of any one of claims 54-64, wherein at least one tonicifying agent is selected from the group comprising: NaCl, KCl, NaNO<sub>3</sub>, Na<sub>2</sub>SO<sub>4</sub>, Na<sub>2</sub>HPO<sub>4</sub>, NaH<sub>2</sub>PO<sub>4</sub>, NaNO<sub>3</sub>, KNO<sub>3</sub>, K<sub>2</sub>SO<sub>4</sub>, K<sub>2</sub>HPO<sub>4</sub>, KH<sub>2</sub>PO<sub>4</sub>, and KNO<sub>3</sub>.  
[0112] 67. The cell culture system of any one of claims 54-64, wherein one tonicifying agent is NaCl.

[0113] 68. The cell culture system of claim 67, wherein the concentration of NaCl in the cell culture medium is 4.5 g/L or higher.  
[0114] 69. The cell culture system of claim 67, wherein the concentration of NaCl in the cell culture medium is 6.5 g/L or higher.  
[0115] 70. The cell culture system of claim 67, wherein the concentration of NaCl in the cell culture medium is 7 g/L or higher.  
[0116] 71. The cell culture system of claim 67, wherein the concentration of NaCl in the cell culture medium is 7.5 g/L or higher.  
[0117] 72. The cell culture system of any one of claims 54-71, wherein the cell culture medium contains an ionic tonicifying agent at a concentration sufficient to produce at least a 50% increase in total rAAV production and a 20% decrease in helper virus production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0118] 73. The cell culture system of claim 72, wherein the cell culture medium contains an ionic tonicifying agent at a concentration sufficient to produce at least a 100% increase in total rAAV production and a 30% decrease in helper virus production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0119] 74. The cell culture system of claim 72, wherein the cell culture medium contains an ionic tonicifying agent at a concentration sufficient to produce at least a 150% increase in total rAAV production and a 40% decrease in helper virus production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0120] 75. The cell culture system of claim 72, wherein the cell culture medium contains an ionic tonicifying agent at a concentration sufficient to produce at least a 200% increase in total rAAV production and a 50% decrease in helper virus production compared to a host cell incubated in a medium with an osmolality of 266 mOsm/kg.  
[0121] 76. The cell culture system of any one of claims 54-64, wherein the tonicifying agent is a non-ionic tonicifying agent.  
[0122] 77. The cell culture system of claim 76, wherein the tonicifying agent is a sugar.  
[0123] 78. The cell culture system of claim 77, wherein the tonicifying agent is a disaccharide.  
[0124] 79. The cell culture system of claim 78, wherein the tonicifying agent is selected from the group consisting of sucrose, fructose, glucose, galactose, mannose, maltose, and trehalose.  
[0125] 80. The cell culture system of claim 79, wherein the tonicifying agent is sucrose.  
[0126] 81. The cell culture system of claim 80, wherein the concentration of sucrose in the cell culture medium is 6.8 g/L or higher.  
[0127] 82. The cell culture system of claim 80, wherein the concentration of sucrose in the cell culture medium is 13.7 g/L or higher.  
[0128] 83. The cell culture system of claim 80, wherein the concentration of sucrose in the cell culture medium is 29.4 g/L or higher.  
[0129] 84. The cell culture system of claim 80, wherein the concentration of sucrose in the cell culture medium is 41.1 g/L or higher.  
[0130] 85. The cell culture system of any one of claims 54-84, wherein the cell culture medium is a serum-free cell culture medium.